



UNITED STATES PATENT AND TRADEMARK OFFICE

UNITED STATES DEPARTMENT OF COMMERCE
United States Patent and Trademark Office
Address: COMMISSIONER FOR PATENTS
P.O. Box 1450
Alexandria, Virginia 22313-1450
www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/035,045	01/03/2002	Jon Elliot Adler	P 280681 2001-019	3276
7590 12/15/2004				
CROWELL & MORING LLP		EXAMINER		
P O BOX 14300		BRANNOCK, MICHAEL T		
WASHINGTON, DC 20044-4300				
		ART UNIT	PAPER NUMBER	
		1646		
DATE MAILED: 12/15/2004				

Please find below and/or attached an Office communication concerning this application or proceeding.

DETAILED ACTION

Status of Application: Claims and Amendments

Claims 120-198, 223-234 are withdrawn from further consideration pursuant to 37 CFR 1.142(b), as being drawn to a nonelected invention, there being no allowable generic or linking claim. Applicant timely traversed the restriction (election) requirement in the reply filed on 9/30/04.

Applicant's election with traverse of Group I, claims 1-199, 132-148, 150, 151, 177-222 as the claims relate to the elected species of hT1R2 is acknowledged. However, the examiner finds that only claims 1-119, 199-222 read on the elected species.

The traversal is on the grounds that polynucleotide claims and encoded polypeptide claims should be examined together and that a search of both would not be a serious burden on the examiner. This is not found persuasive for the following reasons:

Under MPEP § 803, there are two criteria for a proper requirement for restriction between patentably distinct inventions:

(A) The inventions must be independent (see MPEP § 8702.01, 806.04, 808.01) or distinct as claimed (see MPEP § 806.05- §806.05(I)); and

(B) There must be a serious burden on the examiner if restriction is required (see MPEP § 803.02, § 806.04(a)- 806.04(I), § 808.01(a), and § 808.02).

Consistent with current patent practice, a serious search burden may be established by (A) separate classification thereof: (B) a separate status in the art when they are classifiable together: (C) a different field of search. These criteria were met in the above restriction. Further, a search is directed not only to art which would be anticipatory, but also to art that would render the invention obvious. In the instant case, although a search of the polypeptides of Group II would overlap a search of the polynucleotides of Group I, the two searches would not

Art Unit: 1646

be coextensive. In many instances, a protein will have been known in the art before the DNA has been discovered that encodes the protein. Often the protein will be known by a name different than the name given the protein after the cloning of the nucleic acid - and may even be associated with a completely different activity than that ascribed to it when the nucleic acid was cloned. Thus, Groups I and II require divergent searches, and to search both inventions would be burdensome. Therefore, the restriction is maintained and made final

Specification

The disclosure is objected to because it contains an embedded hyperlink and/or other form of browser-executable code, see pages 16 and 39 for example. Applicant is required to delete the embedded hyperlink and/or other form of browser-executable code. See MPEP § 608.01.

Additionally, the disclosure is objected to because the US Provisional Application Number at Page 10 is left blank. Correction is required. See MPEP § 608.01(b).

Claim Rejections - 35 USC § 112

The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

Claims 1-5, 10, 11, 14, 15, 24, 25, 34, 35, 44, 45, 56-58, 68, 69, 80, 81, 90-119 201, 202, 213, 214 are rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

Art Unit: 1646

Claims 1, 90, 103 and dependent claims, require a "variant" of a DNA. The word "variant" is used in the art to denote a relative relationship between two things, yet the specification has not set forth a clear distinction between what is to be considered a "variant" and what is considered to be unrelated. Thus, the artisan could not unambiguously know whether or not he or she was in possession of polynucleotides that are encompassed by Applicants claims. Additionally, claim 103 requires a "variant molecule". The sentence structures of the claims do not explicitly set forth what reference molecule the claimed variant molecule is a variation of, therefore the metes and bounds of the claim cannot be determined.

Claim 2 recites the phrase "consists essential of SEQ ID NO: 15 or 20". This appears to be a minor typographical error, however the phrase renders the claims indefinite because the artisan cannot determine what is and what is not being claimed. For the purpose of this examination, it is assumed that Applicant intended the phrase to read "consisting essentially of".

Claims 4, 5, 14, 15, 24, 25, 34, 35, 44, 45, 56, 57, 68, 69, 80, 81, 92, 93, 105, 106, 201, 202, 213, 214, require that the nucleic acid hybridize under stringent conditions. The term "stringent conditions" is a relative term and encompasses conditions of varying degrees of stringency - such conditions determining the bounds of the claim. However, the art does not provide an unambiguous definition of the term "stringent conditions" and neither is such a definition given for the term in the specification which puts forth the metes and bounds of the claim Applicant is seeking protection for. The term appears to be defined only by way of example at page 30. It is suggested that the claim recite the actual conditions that applicant considers to be stringent, e.g., salt concentration and temperature conditions of incubations and washes.

Art Unit: 1646

Claim 58 requires an isolated fragment “of the genomic DNA molecule of claim 54”, yet there is no genomic DNA molecule of claim 54, thus the artisan would not know which genomic DNA molecule the claim is referring to.

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

Claims 1, 4-11, 14-41, 44-51, 56-119, 201-208, 211-222 rejected under 35 U.S.C. 112, first paragraph, because the specification, while being enabling for isolated naturally occurring polynucleotides that hybridize to a polynucleotide of SEQ ID NO: 20 under the stringent conditions set forth in lines 17 and 18 of page 16 of the specification and encode a polypeptide that bind sucrose in conjunction with a T1R3 polypeptide, does not reasonably provide enablement for artificially constructed polynucleotides that encode variants of the polypeptide of SEQ ID NO: 21, and nor for fragments of the polynucleotide of SEQ ID NO: 20. The specification does not enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the invention commensurate in scope with these claims.

The claims encompass polynucleotides encoding polypeptide variants of the polypeptide of SEQ ID NO: 21, i.e. substitutions, deletions or insertions in a protein corresponding to SEQ ID NO: 21. Applicant has not provided sufficient guidance as to how to make and use the encoded polypeptides which are not 100% identical to the polypeptide of SEQ ID NO: 21, but which still retain a desired property of the polypeptide of SEQ ID NO: 21.

Art Unit: 1646

The specification has failed to teach one of skill in the art which amino acid substitutions, deletions or insertions to make. Furthermore, Applicant has not provided guidance as to what properties of the allelic variants or sequence variants of the protein corresponding to SEQ ID NO: 4 might be desired nor any guidance as to which amino acid substitutions, deletions or insertions to make to achieve any desired property. Applicant has not defined a difference in structure or difference in function between the protein corresponding to SEQ ID NO: 21 and variants of said protein. If a variant of the protein corresponding to SEQ ID NO: 21 is to have a structure and function similar to the protein corresponding to SEQ ID NO: 21 then the specification has failed to teach one of skill in the art which amino acid substitutions, deletions or insertions to make that will preserve the structure and function of the protein corresponding to SEQ ID NO: 21.

The problem of predicting protein structure from sequence data and in turn utilizing predicted structural determinations to ascertain functional aspects of the protein is extremely complex. While it is known that many amino acid substitutions are generally possible in any given protein, the positions within the protein's sequence where such amino acid substitutions can be made with a reasonable expectation of success are limited. Certain positions in the sequence are critical to the protein's structure/function relationship, e.g. such as various sites or regions directly involved in binding, activity and in providing the correct three-dimensional spatial orientation of binding and active sites. These regions can tolerate only relatively conservative substitutions or no substitutions (see Bowie et al., 1990, Science 247:1306-1310, especially p.1306, column 2, paragraph 2). However, Applicant has provided little or no guidance beyond the mere presentation of sequence data to enable one of ordinary skill in the art

Art Unit: 1646

to determine, without undue experimentation, the positions in the protein which are tolerant to change (e.g. such as by amino acid substitutions or deletions), and the nature and extent of changes that can be made in these positions. Also, these or other regions may be critical determinants of antigenicity. It is well appreciated in the art of antibody production that it is unpredictable which amino acids are critical antigenic determinants (see Alexander et al., Proc. Natl. Acad. Sci. 89(3352-3356)1992. Protein antigenicity can be significantly reduced by substitution of even a single residue. Further, even if an amino acid substitution does not destroy the activity of the immunizing protein, the substitution may significantly reduce the antigenicity of the protein (see the Abstract of Alexander et al.). The specification does not provide sufficient guidance as to how to make antibodies that are specific to variants of SEQ ID NO: 21 that can be used for any specific purpose. The specification has not provided guidance as to natural variants that may exist, nor how to use antibodies specific to variants that might be created.

Although the specification outlines art-recognized procedures for producing variants, this is not adequate guidance as to the nature of active variants that may be constructed, but is merely an invitation to the artisan to use the current invention as a starting point for further experimentation. Even if an active or binding site were identified in the specification, they may not be sufficient, as the ordinary artisan would immediately recognize that an active or binding site must assume the proper three-dimensional configuration to be active, which conformation is dependent upon surrounding residues; therefore substitution of non-essential residues can often destroy activity.

The problem of producing active variants appears especially difficult in the art of T1R receptors, to which the instant polypeptide is asserted to belong. The instant specification

Art Unit: 1646

appears to simply suggest to the artisan that art-recognized procedures for screening GPCRs (e.g. pages 32-33, 40 and the example at page 91) are sufficient to identify functional variants of SEQ ID NO: 21. However, Hoon *et al.*, *Cell* 96(541-551)1999, report that "We have attempted to determine the ligand/tastant specificity of TR1 and TR2 using a variety of strategies but have been hampered by the difficulty of functionally expressing these molecules in heterologous systems" see col 1 of page 547. The art regarding T1R receptors, as exemplified by Hoon *et al.*, recognizes the complexity, unpredictability, and non-routine nature of the work involved in trying to assay functional T1R receptors. The instant specification has provided only general guidance to the skilled artisan -such guidance does not supply the artisan with the detailed methods one would need to possess in order to screen for functional variants. Further, the specification has offered no working example of such variants

Additionally, the specification has provide no specific information as to which of multitude of the small fragments of DNA, e.g. claim 6, can be used for any specific purpose.

Therefore, due to the large quantity of experimentation necessary to generate the infinite number of variant recited in the claims and possibly screen same for activity, the lack of direction/guidance presented in the specification regarding which structural features are required in order to provide activity, the absence of working examples directed to same, the complex nature of the invention, the state of the prior art which establishes the unpredictability of the effects of mutation on protein structure and function and the difficulties encountered in screening T1Rs, exemplified by Hoon *et al.*, and the breadth of the claims which fail to recite any structural or functional limitations, undue experimentation would be required of the skilled artisan to make and/or use the claimed invention in its full scope.

Art Unit: 1646

Claims 1, 4, 5, 7-11, 14, 15, 17-41, 44, 45, 47-53, 56, 57, 59-119, 201, 202, 204-208, 211-222 are rejected under 35 U.S.C. 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention.

The specification discloses a naturally occurring polynucleotide of SEQ ID NO: 20 encoding a polypeptide of SEQ ID NO: 21, yet the claims encompass polynucleotides not described in the specification, e.g., artificially mutated sequences, sequences that have a recited degree of identity or that merely hybridize to SEQ ID NO: 20. These claimed genera do not meet the written description provision of 35 U.S.C. 112, first paragraph. Although one of skill in the art would reasonably predict that these sequences exist or could be made to exist, one would not be able make useful predictions as to the nucleotide positions or identities of those sequences based on the information disclosed in the specification.

A genus claim may be supported by a representative number of species as set forth in *Regents of the University of California v Eli Lilly & Co*, 119F3d 1559, 1569, 43 USPQ2d 1398, 1406 (Fed. Cir. 1997). A description of a genus of cDNAs may be achieved by means of a recitation of a representative number of cDNAs, defined by nucleotide sequence, falling within the scope of the genus, or of a recitation of structural features common to the genus, which features constitute a substantial portion of the genus. The instant specification discloses no artificially mutated sequences that have any function. Further, even if the disclose sequence were definitive of a genus with a specified function, the instantly claimed genus is not so limited

Art Unit: 1646

and the prior art does not provide compensatory structural or correlative teachings to enable one of skill to identify/obtain the polynucleotides encompassed, thus the artisan would not consider Applicant to be in possession of the breadth that is claimed.

With the exception of the of the polynucleotide of SEQ ID NO: 20, the skilled artisan cannot envision the detailed chemical structure of the encompassed variants. Therefore, only the polynucleotide of SEQ ID NO: 20, other polynucleotides that encode a polypeptide of SEQ ID NO: 21, and polynucleotides consisting of fragments thereof, but not the full breadth of the claims meet the written description provision of 35 U.S.C. §112, first paragraph.

Claim Rejections - 35 USC § 102

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

(e) the invention was described in (1) an application for patent, published under section 122(b), by another filed in the United States before the invention by the applicant for patent or (2) a patent granted on an application for patent by another filed in the United States before the invention by the applicant for patent, except that an international application filed under the treaty defined in section 351(a) shall have the effects for purposes of this subsection of an application filed in the United States only if the international application designated the United States and was published under Article 21(2) of such treaty in the English language.

Claims 1, 6, 14-16, 22-26, 32-36, 44, 45, 56, 57, 66-70, 76-82, 88-94, 100-107, 113-118, 201-203, 211-215, 221, 222 are rejected under 35 U.S.C. 102(b) as being anticipated by Hoon *et al.*, *Cell* 96(541-551), February 19, 1999.

Art Unit: 1646

Hoon et al. disclose a polynucleotide encoding a polypeptide with over 70% sequence identity with the instant SEQ ID NO: 20, see attached sequence alignment, and would thus be expected to hybridize under what could be considered stringent hybridization conditions as stated in the claims, absent evidence to the contrary. Isolated RNA is also described, page 542. PCR primers are disclosed, page 544, which would meet the conditions of being between about 20-30 nucleotide bases in length, absent evidence to the contrary.

Claims 7-11, 17-21, 27-31, 37-41, 47-51, 59-63, 71-75, 83-87, 95-99, 103-115, 204-208, 216-220 are rejected under 35 U.S.C. 102(b) as being anticipated by Krautwurst et al. Cell 95(917-926)1998.

Claims 7-11, 17-21, 27-31, 37-41, 47-51, 59-63, 71-75, 83-87, 95-99, 108-112, 204-208, 216-220 require that the chimeric or fused nucleic acid molecule only comprise at least a part of the coding sequence contained in the DNA of SEQ ID NO: 20. This limitation reads on any chimeric or fused nucleic because the parts of the coding sequence of SEQ ID NO: 20 are simply A, T, G, or C. Krautwurst et al. disclose a fusion protein comprising a nucleic acid encoding an olfactory receptor (GPCR) fused to a nucleic acid sequence encoding a mammalian rhodopsin (heterologous GPCR), wherein such rhodopsin gene product helps facilitate expression of the fusion protein to the cell surface (pg 918 col 1), wherein the rhodopsin gene product further provides a detectable marker (pg 918 col 2). Such fused nucleic acid is produced with a constitutively active promoter (page 924, col 2). Claims 103-115 and 117 require a variant of SEQ ID NO: 21 containing at least one conservative substitution. Absent evidence to the contrary this reads on any polypeptide.

Art Unit: 1646

Claims 1, 4-7, 9, 11, 14-17, 19, 21-27, 29, 31-37, 39, 41, 44-46, 56-58, 66-171, 73, 75-83, 85, 87-89, 103-115, 117, 201-204, 206, 208-217, 218, 220-222 are rejected under 35 U.S.C. 102(e) as being anticipated by U.S. Patent Application Publication No: 203/0036089, which is fully supported by U.S. Provisional Application No: 60/172,600, filed 12/20/1999

U.S. Patent Application Publication No: 203/0036089 discloses polynucleotides, cDNA, genomic DNA, and expressed RNA [0049], wherein the polynucleotides have 95% identity with the polynucleotide of SEQ ID NO: 20 over the entire length of SEQ ID NO: 20 see attached sequence alignment, and would thus be expected to hybridize to the instant SEQ ID NO: 20 under conditions listed in the specification as stringent, although the polynucleotide is not asserted to be active in taste signaling. Fusion proteins to facilitate purification or expression are also contemplated [0071], as well as probes and primers of about 12-50 nucleotides [0151].

Claim Rejections - 35 USC § 103

The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:

(a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negated by the manner in which the invention was made.

Claims 7-11, 27-31, 37-41, 71-75, 83-87, 95-99, 109-112, 216-220 are rejected under 35 U.S.C. 103(a) as being unpatentable over Hoon *et al.*, *Cell* 96(541-551), February 19, 1999 in view of Krautwurst *et al.* *Cell* 95(917-926)1998.

Art Unit: 1646

As set forth above Hoon et al. disclose a polynucleotide encoding a polypeptide with over 70% sequence identity with the instant SEQ ID NO: 20, see attached sequence alignment, and would thus be expected to hybridize under what could be considered stringent hybridization conditions as stated in the claims, absent evidence to the contrary. Hoon et al. teach that this polynucleotide is difficult to express and explicitly recommend that the expression system of Krautwurst et al. be employed to express the protein, see col 1 of page 547 of Hoon et al.

Krautwurst et al. disclose an expression system producing a fusion protein from a nucleic acid encoding an olfactory receptor (GPCR) fused to a nucleic acid sequence encoding a mammalian rhodopsin (heterologous GPCR), wherein such rhodopsin gene product helps facilitate expression of the fusion protein to the cell surface (pg 918 col 1), wherein the rhodopsin gene product further provides a detectable marker (pg 918 col 2). Such fused nucleic acid is produced with a constitutively active promoter (page 924, col 2).

Therefore, one of ordinary skill in the art, at the time the invention was made, and with reasonable expectation of success, would be motivated to make a chimeric construct with the polynucleotide of Hoon et al. and the rhodopsin construct as taught by Krautwurst et al. The motivation to do so is provided by Hoon et al. who specifically teach to do this.

Allowable Subject Matter

Claims 12, 13, 42, 43, 52-55, 65, 65, 199, 200, 209, 210 are allowable with respect to SEQ ID NO: 20 and 21, yet the claims are objected to because they contain non-elected subject matter.

Art Unit: 1646

Conclusion

Please note the new central fax number for official correspondence below:

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Michael Brannock, Ph.D., whose telephone number is (571) 272-0869. The examiner can normally be reached on Mondays through Fridays from 10:00 a.m. to 4:00 p.m.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Brenda Brumback, Ph.D., can be reached at (571) 272-0961. Official papers filed by fax should be directed to 571-273-8300.

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to the Group receptionist whose telephone number is (703) 308-0196.

MB



December 11, 2004



ELIZABETH KEMMERER
PRIMARY EXAMINER


```

Q9ZOR7
ID Q9ZOR7 PRELIMINARY; PRT; 843 AA.
AC Q9ZOR7;
DT 01-MAY-1999 (TrEMBLrel. 10, Created)
DT 01-MAY-1999 (TrEMBLrel. 10, Last sequence update)
DT 01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DE Putative taste receptor TR2 (Fragment).
OS Rattus norvegicus (Rat).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.
OX NCBI_TaxID=10116;
RN [1]
RP SEQUENCE FROM N.A.
RC STRAIN=Wistar;
RX MEDLINE=99159821; PubMed=10052456;
RA Hoon M.A., Adler E., Lindemeier J., Battey J.F., Ryba N.J.,
RA Zuker C.S.;
RT "Putative mammalian taste receptors: a class of taste-specific GPCRs
RT with distinct topographic selectivity.";
RL Cell 96:541-551(1999).
DR EMBL; AF127390; AAD18070.1; -
DR GO; GO:0016020; C:membrane; IEA.
DR GO; GO:0008067; F:metabotropic glutamate, GABA-B-like recepto. . .; IEA.
DR GO; GO:0004872; F:receptor activity; IEA.
DR InterPro; IPR001828; ANF receptor.
DR InterPro; IPR000337; GPCR_Mgr.
DR InterPro; IPR011500; NCD3G_GPCR.
DR Pfam; PF00003; 7tm_3; 2.
DR Pfam; PF01094; ANF receptor; 1.
DR Pfam; PF07562; NCD3G; 1.
DR PRINTS; PR00248; GPCRMRGR.
DR PROSITE; PS50259; G_PROTEIN_RECEP_F3_4; 1.
KW Receptor.
FT NON_TER 843 843
SQ SEQUENCE 843 AA; 95799 MW; D23AC22D21E049B8 CRC64;

```

Query Match 72.7%; Score 3231; DB 2; Length 843;
Best Local Similarity 70.8%; Pred. No. 3.3e-231;
Matches 596; Conservative 109; Mismatches 133; Indels 4; Gaps 2;

Qy	1	MGPRAKTICSLFFLLWVLAERP---AENSDFYLPGDYLLGGLFSLHANMKGI VHLNFWLQVP	57
Db	1	MGPOARTLCLLSLLHLVLPKPGKLVENSDFHLAGDYLLGGLFTLHANVKSISHLSYLQVP	60
Qy	58	MCKEYEVKIVIGYNLMQAMRF AVEEINNDSLLPGVLLGYEIVDVCYISNNVQPVLYFLAH	117
Db	61	KCNEFTMKVLGYNLMQAMRF AVEEINNCSLLPGVLLGYEMVDVCYLSNNIHGPLYFLAQ	120
Qy	118	EDNLLPIQEDYSNYISRVVAVIGPDNSESVMTVANFLSLFLLPQITYSAISDEL RDKVRF	177
Db	121	DDDLLPILKDYSQYMPHVAVIGPDNSESAITVSNILSHFLIPQITYSAISDKLRDKRHF	180
Qy	178	PALLRTTPSADHHVEAMVQLMLHFRWNWIIVLVSSDITYGRDNGQLGERVAR-RDICIAF	236
Db	181	PSMLRTVPSATHHIEAMVQLMVHPQWNWIIVLVSDDDYGRENSHLLSQRLTXTSDICIAF	240
Qy	237	QETLPTLQPNQNM TSEERQRLVTIVDKLQQSTARVVVVFSPDLTYHFFNEVLRQNFTGA	296
Db	241	QEVLPPIESSQVMRSEEQRLDNILDKLRRTSARVVVVFSPELSLYSFFHEVLRWNFTGF	300
Qy	297	VWIASESWAIDPVLHNLTELGH LGTFLGITIQSVPIPGFSEFREWGPQAGPPPLSRTSQS	356
Db	301	VWIASESWAIDPVLHNLTEL RHTGTF LGVTIQRVSI PGFSQFRVRDKPGYPV PNTTNLR	360
Qy	357	YTCNQECDNCLNATLSFNTILRLSGERVVYSVYS AVYAVAHALHSLGCDKSTCTKRVVY	416
Db	361	TTCNQDCDACLN TTKSFNNILILSGERVVYSVYS AVYAVAHALHRLG CNRVRCTKQKVY	420
Qy	417	PWOLLEEIWKNFTLLDHQIFFDPOGDVALHLEIVQWQWDRSQNP FQSVASYPLQRQLK	476
Db	421	PWQLLREIWHVNFTLLGNRLFDDQQGDMPMLLDIIQWQWDL SQNP FQSIASYSPTSKRLT	480
Qy	477	NIQDISWHTVNNTIPMSMCKSRCSQGQKKPVGIHVCCFECIDCLPGTFLNHTED EYECQ	536
Db	481	YINNVSWYTPNNTVPVSMCKSKCQPGQMKS SVGLHPCCFECLDCMPGTYLNRSADEFNCL	540
Qy	537	ACPNNWSYQSETSCFKRQLVFLEWHEAPTIAVALLAALGFLSTLAILVIFWRHFQTPIV	596
Db	541	SCPGSMWSYKNDITCFQRRPTFLEWHEVPTIVVAILAALGFFSTLAILFIFWRHFQTPMV	600
Qy	597	RSAGGPMCFLMLTLLLVA YMVVPVYVGPPKVSTCLCRQALFPLCFTICISCI AVR SFQIV	656
Db	601	RSAGGPMCFLMLVPLLLAFGMVVPVYGPPTV FSCFCRQAFFTVCFSICLSCITVRSFQIV	660
Qy	657	CAFKMASRFPRAYSYWVRYQGPYVSMAFITVLK MVI VVIGMLATGLSPTTRTDPDDPKIT	716
Db	661	CVFKMARLEPSAYS FWMRYHGPYVFVAFITA I KVALVVG NMLATTINPIGRTPDDPNIM	720
Qy	717	IVSCNPNYRNSLLFN TSLD LLSVVGFSFAYMGKELPTNYNEAKFITLSMTFYFTSSVSL	776
Db	721	ILSCHPNYRNGLLFN TSM D LLSVLGFSFAYMGKELPTNYNEAKFITLSMTFSFTSSISL	780
Qy	777	CTFMSAYSGVLVTIVDLLVTVLNLLAISLGYFGPKCYMILFYPERNTPAYFN SMIOGYTM	836
Db	781	CTFMSVHDGVLVTIMDLLVTVLNFLAIGLGYFGPKCYMILFYPERNTSAYFN SMIOGYTM	840

```

RESULT 8
US-10-261-482-3
; Sequence 3, Application US/10261482
; Publication No. US20030036089A1
; GENERAL INFORMATION:
; APPLICANT: WEI, Ming-Hui et al
; TITLE OF INVENTION: ISOLATED HUMAN G-PROTEIN COUPLED
; TITLE OF INVENTION: RECEPTORS, NUCLEIC ACID MOLECULES ENCODING HUMAN GPCR
; TITLE OF INVENTION: PROTEINS, AND USES THEREOF
; FILE REFERENCE: C1000869CON
; CURRENT APPLICATION NUMBER: US/10/261,482
; CURRENT FILING DATE: 2002-10-02
; PRIOR APPLICATION NUMBER: 09/684,393
; PRIOR FILING DATE: 2000-10-10
; PRIOR APPLICATION NUMBER: 60/172,600
; PRIOR FILING DATE: 1999-12-20
; NUMBER OF SEQ ID NOS: 10
; SOFTWARE: FastSeq for Windows Version 4.0
; SEQ ID NO 3
; LENGTH: 8001
; TYPE: DNA
; ORGANISM: Human
US-10-261-482-3.

```

Application/Control Number: 10/035,045

Query Match	95.5%	Score 3403.8	DB 14	Length 8001
Best Local Similarity	97.9%	Pred. No. 0		
Matches 3489	Conservative	0	Mismatches 22	Indels 52
				Gaps 22
QY	1	AGCCTGGCAGTGGCCCTCAGGCAAGTCTGACGCGCACAACTTTACAGGCCAGGAAGCGA	60	
Db	1908	AGCCTGGCAGTGGCCCTCAGGCAAGTCTGACGCGCACAACTTTACAGGCCAGGAAGCGA	1967	
QY	61	GGAACCACTGGGGCCCCAGGGTGTGGCAAGTGAAGATGGCAAGGTTTGTCTAAACAA	120	
Db	1968	GGAACCACTGGGGCCCCAGGGTGTGGCAAGTGAAGATGGCAAGGTTTGTCTAAACAA	2027	
QY	121	TCCTCTGCCCCGCTCCCCCGCCCCGGGCTCACTCCATGTGAAGCCCCCATCGGGGCAAGCCAC	180	
Db	2028	TCCTCTGCCCCGCTCCCCCGCCCCGGGCTCACTCCATGTGAAGCCCCCATCGGGGCAAGCCAC	2087	
QY	181	CTGCCGTGCTGTGTGAAGTTGCTCTGCCATGCTGGGCCCTGTGCTCTGGGCTCTAGC	240	
Db	2088	CTGCCGTGCTGTGTGAAGTTGCTCTGCCATGCTGGGCCCTGTGCTCTGGGCTCTAGC	2147	
QY	241	CTCTGGGCTCTCCTGCAACCTTGGGACGGGGGCCCATTTGTGCTGTCAACGCACTTAGG	300	
Db	2148	CTCTGGGCTCTCCTGCAACCTTGGGACGGGGGCCCATTTGTGCTGTCAACGCACTTAGG	2207	
QY	301	ATGAAGGGGACTACGTGCTGGGGGGGCTGTTCCCCCTGGGCGAAGCCGAGAGGCTGGC	360	
Db	2208	ATGAAGGGGACTACGTGCTGGGGGGGCTGTTCCCCCTGGGCGAAGCCGAGAGGCTGGC	2267	
QY	361	CTCCGAGCCGGAACAGGCCACAGCCCTGTGTGCAACAGTACAGAGGTGGACGGCC	420	
Db	2268	CTCCGAGCCGGAACAGGCCACAGCCCTGTGTGCAACAGTACAGAGGTGGACGGCC	2327	
QY	421	TGGGTGGGGTCAAGGTGACAGGTCTGGGGTGCTCCTGAGCTGGGGCCGAGGTGGCCAT	480	
Db	2328	TGGGTGGGGTCAAGGTGACAGGTCTGGGGTGCTCCTGAGCTGGGGCCGAGGTGGCCAT	2387	
QY	481	CTGCGGTTCTGTGTGGCCCAAGTTCTCCTCAAAAGGCGCTGCTGGGCACTGGCCATGA	540	
Db	2388	CTGCGGTTCTGTGTGGCCCAAGTTCTCCTCAAAAGGCGCTGCTGGGCACTGGCCATGA	2447	
QY	541	AAATGGCCGTGAGAGATCAACAAGTCGATCTGTGCCCGGGCTGCGCTGGGCT	600	
Db	2448	AAATGGCCGTGAGAGATCAACAAGTCGATCTGTGCCCGGGCTGCGCTGGGCT	2507	

QY 601 ACCAGCCTTTGATACGTCCTCGGAGCCTGTGGTGGCCATGAAGCCCAAGCTCATGTTCC
 Db 2508 ACGACCTCTTTGATACGTGCTCGGAGCCTGTGGTGGCCATGAAGCCCAAGCTCATGTTCC
 QY 661 TGGCCAAAGGCAGGCAAGCCGCGACATCGCCGCTACTGCAACTACACGCAGTACAGCCCCC
 Db 2568 TGCGCCAAAGGCAGGCAAGCCGCGACATCGCCGCTACTGCAACTACACGCAGTACAGCCCCC
 QY 721 GTGTGCTGGCTGTCAATCGGGGCCCACTCGTCAAGAGTCGCCATGTCACCCGCAAGTTCT
 Db 2628 GTGTGCTGGCTGTCAATCGGGGCCCACTCGTCAAGAGTCGCCATGTCACCCGCAAGTTCT
 QY 781 TCAGCTTCTTCTCATGCCCCCAGTGGGGCGCCCCCAACCATCACCCCAACCCCAACCAACC
 Db 2688 TCAGCTTCTTCTCATGCCCCCAGTGGGGCGCCCCCAACCATCACCCCAACCCCAACCAAGCC
 QY 841 CCTGCCCCGTGGGAGCCCTTGTGTCAAGAGATGCTACATGCACCCCAACCAAGCCCTGC
 Db 2748 CTGCCCCGTGGAG--CCCTGTGTCAAGAGATGCT-----
 QY 901 CCTGGGAAGCCCTGTGTCAAGAGATGCTCTTTGGCCTTGCAGGTCAAGCTACGGTGTAGCAT
 Db 2782 -----CCTGGCCTTGCAGGTCAAGCTACGGTGTAGCAT
 QY 961 GGAAGTGTCTGAGCGCCCGGAGAGACCTTCCCTCTCTTCTTCCGCAACCGTGGCCAGCGACCG
 Db 2816 GGAAGTGTCTGAGCGCCCGGAGAGACCTTCCCTCTCTTCTTCCGCAACCGTGGCCAGCGACCG
 QY 1021 TGTGCAGCTGACGCGCCGCGCGGAGAGCTGCTGCAGAGAGTTCCGGCTGGAAGTACTGGGTGGCCGC
 Db 2876 TGTGCAGCTGACGCGCCGCGCGGAGAGCTGCTGCAGAGAGTTCCGGCTGGAAGTACTGGGTGGCCGC
 QY 1081 CCTGGGCAAGCGACGACGAGTACGGCCGCGCAAGGAGCTTGAAGCATCTTCTCGGCTTGGCCGC
 Db 2936 CCTGGGCAAGCGACGAGTACGGCCGCGCAAGGAGCTTGAAGCATCTTCTCGGCTTGGCCGC
 QY 1141 GGCACGCGGCGCATCTGCATTCGCGCAAGAGGCTGGTCCGCTGCCCGGTGCCATGACTC
 Db 2996 GGCACGCGGCGCATCTGCATTCGCGCAAGAGGCTGGTCCGCTGCCCGGTGCCATGACTC
 QY 1201 GCGGCTGGGGAAGGTGCAGGACGTCTCTGCACCAAGTGAACCAAGACAGCGTGCAGGTGGT
 Db 3056 GCGGCTGGGGAAGGTGCAGGACGTCTCTGCACCAAGTGAACCAAGACAGCGTGCAGGTGGT
 QY 1261 GCTGCTGTTGCGCTCCGTGCACGCGCGCCCAAGCCTCTTCAACTACAGCATCAGCAGCAG
 Db 3116 GCTGCTGTTGCGCTCCGTGCACGCGCGCCCAAGCCTCTTCAACTACAGCATCAGCAGCAG
 QY 1321 GCTCTCGCCCCAAGGTGTGGGTGGCCAGCGAGGCTGAGCCTCTGACCTGTGCATGGG
 Db 3176 GCTCTCGCCCCAAGGTGTGGGTGGCCAGCGAGGCTGAGCCTCTGACCTGTGCATGGG
 QY 1381 GCTGCCCCGCGCATGCGCCAGATGGGCAAGTGTGGCTTCTCTCAAGAGGGGTGCCAGCT
 Db 3236 GCTGCCCCGCGCATGCGCCAGATGGGCAAGTGTGGCTTCTCTCAAGAGGGGTGCCAGCT
 QY 1441 GCAAGAGTTCCTCCCACTACGTGAAGACGCACTTGGCCCTGSCACCGAACCCGSCCTTCTG
 Db 3296 GCAAGAGTTCCTCCCACTACGTGAAGACGCACTTGGCCCTGSCACCGAACCCGSCCTTCTG
 QY 1501 CTCTGCCCCGTGGCGAAGGGAGCAGGCTGTGAAGAGAGACGTGGTGGGCGCAGCGCTGCC
 Db 3356 CTCTGCCCCGTGGCGAAGGGAGCAGGCTGTGAAGAGAGAGACGTGGTGGGCGCAGCGCTGCC
 QY 1561 GCAAGTGTGACTGCATCAGCGTGCAGAACGTGAGCGCAGGCTAAATCACCAACAGACGTT
 Db 3416 GCAAGTGTGACTGCATCAGCGTGCAGAACGTGAGCGCAGGCTAAATCACCAACAGACGTT
 QY 1621 CTCTGTCTACGCAAGCTGTGTATAGCGTGGCCAGGCTGTGACAACTCTTCAGTGCA
 Db 3476 CTCTGTCTACGCAAGCTGTGTATAGCGTGGCCAGGCTGTGACAACTCTTCAGTGCA

Art Unit: 1646

1681 CGCCTCAGGCTGCCCCGCGCAGAGACCCCTGGAAGCCCTGGCAGGTGAGCCCGGAGATGG 1740
1741 GGGTGTGCTGCTCTGATGTGCCCCAGGCCACAGGACAGGCCACCAAGCCTGAGCTGG 1800
3536 CGCCTCAGGCTGCCCCGCGCAGAGACCCCTGGAAGCCCTGGCAGGTGAGCCCGGAGATGG 3595
3596 GGGTGTGCTGCTCTGATGTGCCCCAGGCCACAGGACAGGCCACCAAGCCTGAGCTGG 3655
1801 AGGTGCTGAGGCTCAGCCCCGCTGCCCCGCGCAGCTCCTGGAAGACATGTAACAACCT 1860
3656 AGGTGCTGAGGCTCAGCCCCGCTGCCCCGCGCAGCTCCTGGAAGACATGTAACAACCT 3715
1861 GACCTTCCAGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGG 1920
3716 GACCTTCCAGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGG 3775
1921 GTACGACCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAG 3835
1981 GTTCAACGCGCAGCTCAGAGCAGAGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCT 2040
3836 GTTCAACGCGCAGCTCAGAGCAGAGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCT 3895
2041 GGTGAGGTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAG 2100
3896 GGTGAGGTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAG 3955
2101 CTGGGGGTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAG 2160
3956 CTGGGGGTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAGGCTGAG 4015
2161 CAGGCTGTGCGCAGAGAGCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGT 2220
4016 CAGGCTGTGCGCAGAGAGCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGT 4075
2221 CGCCGGGTCAAGGGGTTCACACTCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 2280
4076 CGCCGGGTCAAGGGGTTCACACTCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4135
2281 TACCGGCAAAACCAAGGTGAGCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGT 2340
4136 TACCGGCAAAACCAAGGTGAGCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGT 4195
2341 AGGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 2400
4196 AGGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4255
2401 GCGCCCTTCT 2460
4256 GCGCCCTTCT 4315
2461 TCCCGGAGGAGGAGCAGAGCAGGCTGCTCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGT 2520
4316 TCCCGGAGGAGGAGCAGAGCAGGCTGCTCCCGGTGCTCCCGGTGCTCCCGGTGCTCCCGGT 4375
2521 CCGGCTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 2580
4376 CCGGCTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4435
2581 TTGGGGCTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTG 2640
4436 TTGGGGCTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTG 4495
2641 GCTGCTTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTG 2700
4496 GCTGCTTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTG 4555
2701 CAGCCCAAGCCCTGCGCATGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 2760
4556 CAGCCCAAGCCCTGCGCATGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4615
2761 TGCCTGAGCACACTTCTCTGAGGCGCGGAGATCTTCTGAGATCAAGAACTGCTCTG 2820

Db 4616 TGCCTGAGCACACTTCTCTGAGGCGCGGAGATCTTCTGAGATCAAGAACTGCTCTG 4675
Qy 2821 AGCTGGGAGACCGGCTGAGTGTGCTGCTGCGGGGCGCTGGGCTGCTGCTGCTGCTGCTG 2880
Db 4676 AGCTGGGAGACCGGCTGAGTGTGCTGCTGCGGGGCGCTGGGCTGCTGCTGCTGCTGCTG 4735
Qy 2881 CTGGCCATGCTGCTGAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 2940
Db 4736 CTGGCCATGCTGCTGAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4795
Qy 2941 CTGGTACGAGCTGGGCTGAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3000
Db 4796 GTGTGACGAGCTGGGCTGAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4855
Qy 3001 TGGGTACGAGCTGGGCTGAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3060
Db 4856 TGGGTACGAGCTGGGCTGAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4915
Qy 3061 GGCACCTTCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3120
Db 4916 GGCACCTTCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 4975
Qy 3121 GCCATGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3180
Db 4976 GCCATGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5035
Qy 3181 GTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3240
Db 5036 GTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5095
Qy 3241 GTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3300
Db 5096 GTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5155
Qy 3301 GAGTTCCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3360
Db 5156 GAGTTCCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5215
Qy 3361 AATCAGGGGAAACATGAGTGAACCAACCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3420
Db 5216 AATCAGGGGAAACATGAGTGAACCAACCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5275
Qy 3421 GCTGCGATCCCCCAAGCAGCAATGACCCGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3480
Db 5276 GCTGCGATCCCCCAAGCAGCAATGACCCGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5335
Qy 3481 AGTTCTGACCCCAAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 3540
Db 5336 AGTTCTGACCCCAAGTGTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCTGCT 5395
Qy 3541 ACCTGACACCCCTGTGACCATC 3563
Db 5396 ACCTGACACCCCTGTGACCATC 5418